10/714,078 Search Lycook 1/12/07

## (FILE 'HOME' ENTERED AT 11:55:10 ON 12 JAN 2007)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, JAPIO' ENTERED AT 11:55:45 ON 12 JAN 2007

	0111 2007	
L1	0	S (108 AMINO ACID BNP)
L2	18	S BNP AND (108 AMINO ACID)
L3	7	DUPLICATE REMOVE L2 (11 DUPLICATES REMOVED)
L4	1	S L3 AND ANTIBOD?
L5	0	S BNP AND NESTRITIDE?
L6	11945	S BNP
L7	330	S L6 AND ANTIBOD?
L8	196	DUPLICATE REMOVE L7 (134 DUPLICATES REMOVED)
L9	6	S L8 AND STROKE?
L10	55	S L8 AND PD<2000
T.11	4.0	S 1.10 AND RPAIN?

```
ANSWER 30 OF 40 EMBASE COPYRIGHT (c) 2007 Elsevier B.V. All rights
     reserved on STN
ΑN
     1998012811 EMBASE.
     Biochemical detection of left-ventricular systolic dysfunction.
TI
     McDonagh T.A.; Robb S.D.; Murdoch D.R.; Morton J.J.; Ford I.; Morrison
ΑU
     C.E.; Tunstall-Pedoe H.; McMurray J.J.V.; Dargie H.J.
     Dr. T.A. McDonagh, Cardiology Department, Western Infirmary, Glasgow G11
CS
     6NT, United Kingdom
so
     Lancet, (3 Jan 1998) Vol. 351, No. 9095, pp. 9-13. .
     Refs: 27
     ISSN: 0140-6736 CODEN: LANCAO
CY
     United Kingdom
     Journal; Article
DT
FS
             Internal Medicine
             Cardiovascular Diseases and Cardiovascular Surgery
     029
             Clinical Biochemistry
             Health Policy, Economics and Management
     036
LA
     English
     English
ED
     Entered STN: 22 Jan 1998
     Last Updated on STN: 22 Jan 1998
     Background: In previous studies on the use of natriuretic peptides to
AB
     detect left-ventricular systolic dysfunction, a higher rate of cardiac
     disorders in the control groups than in the study groups could have led to
           We investigated the effectiveness of plasma N-terminal atrial
     natriuretic peptide (NT-ANP) and brain natriuretic peptide (
     BNP) concentrations to show left-ventricular systolic dysfunction
     in a random sample of the general population. Methods: We randomly
     selected 2000 participants aged 25-74 years from family physicians' lists
     in Glasgow, UK. We sent all participants questionnaires. 1653 respondents
     underwent echocardiography and electrocardiography. We took a
     left-ventricular ejection fraction of 30% or less to show left-ventricular
     systolic dysfunction. NT-ANP and BNP were measured in plasma by
           Findings: 1252 participants had analysable electrocardiograms and
     echocardiograms, completed questionnaires, and available blood samples.
    Median concentrations of NT-ANP and BNP were significantly
    higher in participants with left-ventricular systolic dysfunction (2.8
    ng/mL [IQR 1.8-4.6] and 24.0 pg/mL [18.0-33.0]) than in those without (1.3
    ng/mL [0.9-1.8] and 7.7 pg/mL [3.4-13.0]; each p < 0.001). Among
    participants with left-ventricular systolic dysfunction, both symptomatic
    and asymptomatic subgroups had raised NT-ANP and BNP
     concentrations. A BNP concentration of 17.9 pg/mL or more gave
    a sensitivity of 77% and specificity of 87% in all participants, and 92%
     and 72% in participants aged 55 years or older. Interpretation:
    Measurement of BNP could be a cost-effective method of screening
     for left-ventricular systolic dysfunction in the general population,
     especially if its use were targeted to individuals at high risk.
    Medical Descriptors:
     *heart left ventricle failure: DI, diagnosis
    biochemistry
    hormone blood level
    united kingdom
    echocardiography
    electrocardiography
    heart ejection fraction
    radioimmunoassay
    questionnaire
    cost effectiveness analysis
    high risk population
    human
    male
    female
    major clinical study
    controlled study
```

aged adult clinical trial randomized controlled trial article priority journal Drug Descriptors: \*brain natriuretic peptide: EC, endogenous compound \*atrial natriuretic factor: EC, endogenous compound antibody (brain natriuretic peptide) 114471-18-0; (atrial natriuretic RNfactor) 85637-73-6 (1) RAS 9129; (2) BNP RIK 9086 NP

(2) Peninsula (United States) CO

aged adult clinical trial randomized controlled trial article priority journal Drug Descriptors: \*brain natriuretic peptide: EC, endogenous compound \*atrial natriuretic factor: EC, endogenous compound antibody (brain natriuretic peptide) 114471-18-0; (atrial natriuretic RNfactor) 85637-73-6 NP (1) RAS 9129; (2) BNP RIK 9086

(2) Peninsula (United States) CO

# (FILE 'HOME' ENTERED AT 11:55:10 ON 12 JAN 2007)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, JAPIO' ENTERED AT 11:55:45 ON 12 JAN 2007

L1	0	S (108 AMINO ACID BNP)
L2	18	S BNP AND (108 AMINO ACID)
L3	7	DUPLICATE REMOVE L2 (11 DUPLICATES REMOVED)
L4	1	S L3 AND ANTIBOD?
L5	0	S BNP AND NESTRITIDE?
L6	11945	S BNP
L7	330	S L6 AND ANTIBOD?
L8	196	DUPLICATE REMOVE L7 (134 DUPLICATES REMOVED)
L9	6	S L8 AND STROKE?
L10	55	S L8 AND PD<2000
T.11	4.0	S I.10 AND BRAIN?

RL: ANT (Analyte); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses) (immunoassay with  $\alpha$ - brain natriuretic peptide-specific antibody and prepro-BNP/ $\gamma$ - BNP -specific antibody for BNP determination and cardiac diseases diagnosis) IT 114471-18-0, Brain natriuretic peptide 122007-25-4, Brain natriuretic peptide, prepro-RL: ANT (Analyte); BSU (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses) (immunoassay with  $\alpha$ - brain natriuretic peptide-specific antibody and prepro-BNP/ $\gamma$ - BNP -specific antibody for BNP determination and cardiac diseases diagnosis) IT 124586-56-7 221266-50-8 RL: BSU (Biological study, unclassified); BIOL (Biological study) (immunoassay with  $\alpha$ - brain natriuretic peptide-specific antibody and prepro-BNP/ $\gamma$ - BNP -specific antibody for BNP determination and cardiac diseases diagnosis) THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT

- (1) Anon; FEBS LETTERS 1997, V400(2), P177
- (2) Medinnova Sf; WO 9324531 A CAPLUS
- (3) Medinnova Sf; JP 07507210 A 1995
- (4) Shionogi & Co Ltd; JP 03297392 A 1991 CAPLUS

```
RL: ANT (Analyte); BSU (Biological study, unclassified); PRP (Properties);
     THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study);
     USES (Uses)
        (immunoassay with \alpha- brain natriuretic peptide-specific
        antibody and prepro-BNP/\gamma- BNP
        -specific antibody for BNP determination and cardiac
        diseases diagnosis)
IT
     114471-18-0, Brain natriuretic peptide
                                               122007-25-4,
     Brain natriuretic peptide, prepro-
     RL: ANT (Analyte); BSU (Biological study, unclassified); THU (Therapeutic
     use); ANST (Analytical study); BIOL (Biological study); USES (Uses)
        (immunoassay with \alpha- brain natriuretic peptide-specific
        antibody and prepro-BNP/\gamma- BNP
        -specific antibody for BNP determination and cardiac
        diseases diagnosis)
IT
     124586-56-7
                   221266-50-8
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (immunoassay with \alpha- brain natriuretic peptide-specific
        antibody and prepro-BNP/\gamma- BNP
        -specific antibody for BNP determination and cardiac
        diseases diagnosis)
RE.CNT
              THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
(1) Anon; FEBS LETTERS 1997, V400(2), P177
```

- (2) Medinnova Sf; WO 9324531 A CAPLUS
- (3) Medinnova Sf; JP 07507210 A 1995
- (4) Shionogi & Co Ltd; JP 03297392 A 1991 CAPLUS

```
G01N033/53; G01N033/74
                  ECLA
                         G01N0033-53 [ICS,4]; G01N0033-68 [ICS,4]
 ES 2256952
                  IPCI
                         G01N0033-53 [I,C*]; G01N0033-53 [I,A]; G01N0033-74
                  IPCR
                         [I,C*]; G01N0033-74 [I,A]
                  ECLA
                         G01N033/53; G01N033/74
 NO 2000001273
                  IPCI
                         G01N0033-53 [ICM, 7]
                         G01N0033-53 [I,C*]; G01N0033-53 [I,A]; G01N0033-74
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                         G01N0033-53 [ICM, 7]; C07H0021-04 [ICS, 7]; C07H0021-00
                         [ICS,7,C*]; C12P0021-02 [ICS,7]; C12N0005-06 [ICS,7];
                         C07K0014-47 [ICS,7]; C07K0014-435 [ICS,7,C*]
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                         435/069.100; 435/007.100; 435/320.100; 435/325.000;
                  NCL
                         530/350.000; 536/023.100
                         G01N033/53; G01N033/74
     An immunoassay method specific for mammalian \gamma- BNP derivs.
     which comprises using a first antibody reacting with mammalian
     \alpha- BNP and a second antibody reacting with
     prepro-BNP or y- BNP derivs. but not with
     \alpha- BNP and wherein at least one of these antibodies
     has been detectably labeled or supported on a solid phase.
     immunoassay kit is useful for diagnosis of BNP-associated heart
     diseases.
ST
     monoclonal antibody gamma BNP heart disease
IT
     Blood plasma
     Chemiluminescent substances
     Fluorescent substances
     Heart, disease
     Immunoassay
     Labels
     Mammal (Mammalia)
     Particles
     Test kits
        (immunoassay with \alpha- brain natriuretic peptide-specific
        antibody and prepro-BNP/\gamma- BNP
        -specific antibody for BNP determination and cardiac
        diseases diagnosis)
IT
     Enzymes, biological studies
     Radionuclides, biological studies
     RL: ARU (Analytical role, unclassified); THU (Therapeutic use); ANST
     (Analytical study); BIOL (Biological study); USES (Uses)
        (immunoassay with \alpha- brain natriuretic peptide-specific
        antibody and prepro-BNP/\gamma- BNP
        -specific antibody for BNP determination and cardiac
        diseases diagnosis)
TΤ
     Antibodies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (immunoassay with \alpha- brain natriuretic peptide-specific
        antibody and prepro-BNP/\gamma- BNP
        -specific antibody for BNP determination and cardiac
        diseases diagnosis)
IT
     Immunoassay
        (immunoradiometric assay, sandwich; immunoassay with \alpha-
        brain natriuretic peptide-specific antibody and
        prepro-BNP/\gamma- BNP-specific antibody
        for BNP determination and cardiac diseases diagnosis)
IT
     Antibodies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (monoclonal; immunoassay with \alpha- brain natriuretic
        peptide-specific antibody and prepro-BNP/\gamma-
        BNP-specific antibody for BNP determination and
        cardiac diseases diagnosis)
IT
     121128-24-3, \gamma Brain natriuretic peptide
```

```
ECLA
                         G01N033/53; G01N033/74
                         G01N0033-53 [ICS,4]; G01N0033-68 [ICS,4]
 ES 2256952
                  IPCI
                         G01N0033-53 [I,C*]; G01N0033-53 [I,A]; G01N0033-74
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                         G01N033/53; G01N033/74
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                         G01N0033-53 [ICM, 7]
                         G01N0033-53 [I,C*]; G01N0033-53 [I,A]; G01N0033-74
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                         C07K0014-47 [ICS,7]; C07K0014-435 [ICS,7,C*]
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                         435/069.100; 435/007.100; 435/320.100; 435/325.000;
                         530/350.000; 536/023.100
                  ECLA
                         G01N033/53; G01N033/74
AB
     An immunoassay method specific for mammalian \gamma- BNP derivs.
     which comprises using a first antibody reacting with mammalian
     \alpha- BNP and a second antibody reacting with
     prepro-BNP or \gamma- BNP derivs. but not with
     \alpha- BNP and wherein at least one of these antibodies
     has been detectably labeled or supported on a solid phase.
     immunoassay kit is useful for diagnosis of BNP-associated heart
     diseases.
ST
     monoclonal antibody gamma BNP heart disease
IT
     Blood plasma
     Chemiluminescent substances
     Fluorescent substances
     Heart, disease
     Immunoassay
     Labels
     Mammal (Mammalia)
     Particles
     Test kits
        (immunoassay with \alpha- brain natriuretic peptide-specific
        antibody and prepro-BNP/\gamma- BNP
        -specific antibody for BNP determination and cardiac
        diseases diagnosis)
IT
     Enzymes, biological studies
     Radionuclides, biological studies
     RL: ARU (Analytical role, unclassified); THU (Therapeutic use); ANST
     (Analytical study); BIOL (Biological study); USES (Uses)
        (immunoassay with \alpha- brain natriuretic peptide-specific
        antibody and prepro-BNP/\gamma- BNP
        -specific antibody for BNP determination and cardiac
        diseases diagnosis)
IT
     Antibodies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (immunoassay with \alpha- brain natriuretic peptide-specific
        antibody and prepro-BNP/\gamma- BNP
        -specific antibody for BNP determination and cardiac
        diseases diagnosis)
IT
     Immunoassay
        (immunoradiometric assay, sandwich; immunoassay with \alpha-
        brain natriuretic peptide-specific antibody and
        prepro-BNP/\gamma- BNP-specific antibody
        for BNP determination and cardiac diseases diagnosis)
IT
     Antibodies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (monoclonal; immunoassay with \alpha- brain natriuretic
        peptide-specific antibody and prepro-BNP/\gamma-
        BNP-specific antibody for BNP determination and
        cardiac diseases diagnosis)
IT
     121128-24-3, γ Brain natriuretic peptide
```

```
AN
     1999:194337 CAPLUS
DN
     130:232845
     Entered STN: 25 Mar 1999
ED
     Immunoassay method for brain natriuretic peptide (BNP)
TT
     Asada, Hidehisa; Shimizu, Hiroyuki; Endou, Kazuaki
IN
PA
     Shionogi & Co., Ltd., Japan
SO
     PCT Int. Appl., 24 pp.
     CODEN: PIXXD2
DT
     Patent
LΑ
     Japanese
IC
     ICM G01N033-53
CC
     2-1 (Mammalian Hormones)
     Section cross-reference(s): 9, 15
FAN.CNT 1
     PATENT NO.
                                                                  DATE
                        KIND
                                DATE
                                           APPLICATION NO.
                                            ______
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                        ----
                                ____
                                                                   -----
                                           WO 1998-JP4063
                                19990318
                                                                   19980910 <--
ΡI
     WO 9913331
                         A1
        W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
             DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG,
             KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO,
             NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA,
             UG, US, UZ, VN, YU, ZW
         RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
             FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
             CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     CA 2304263
                         A1
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                                            AU 1998-90010
                                                                   19980910 <--
     AU 731858
                          B2
                                20010405
     EP 1016867
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                                            EP 1998-941797
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     EP 1016867
                          В1
                                20060104
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI, CY
                          B2
                                20040405
                                            JP 2000-511064
     JP 3516655
                                                                   19980910
                          Т
                                            AT 1998-941797
     AT 315228
                                20060215
                                                                   19980910
     ES 2256952
                         Т3
                                20060716
                                           ES 1998-941797
                                                                   19980910
                                           NO 2000-1273
     NO 2000001273
                         Α
                                20000510
                                                                   20000310
                                           US 2000-508435
     US 2003157596
                         A1
                                20030821
                                                                   20000313
                         B2
     US 6828107
                                20041207
PRAI JP 1997-246684
                                19970911
                         Α
     WO 1998-JP4063
                         W
                                19980910
CLASS
PATENT NO.
                CLASS PATENT FAMILY CLASSIFICATION CODES
                ----
                        ------
 -----------
 WO 9913331
                ICM
                       G01N033-53
                        G01N0033-53 [ICM, 6]
                IPCI
                IPCR
                        G01N0033-53 [I,C*]; G01N0033-53 [I,A]; G01N0033-74
                        [I,C*]; G01N0033-74 [I,A]
                ECLA
                        G01N033/53
CA 2304263
                IPCI
                        G01N0033-53 [ICM, 7]
                        G01N0033-53 [I,C*]; G01N0033-53 [I,A]; G01N0033-74
                IPCR
                        [I,C*]; G01N0033-74 [I,A]
AU 9890010
                IPCI
                        G01N0033-53 [ICM, 6]
                        G01N0033-53 [I,C*]; G01N0033-53 [I,A]; G01N0033-74
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EP 1016867
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                        [I,A]; G01N0033-68 [I,A]
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                       G01N0033-53 [I,C*]; G01N0033-53 [I,A]; G01N0033-74
                        [I,C*]; G01N0033-74 [I,A]
                       G01N033/53; G01N033/74
                ECLA
                       G01N0033-53 [ICM, 7]
JP 3516655
                IPCI
                       G01N0033-53 [ICS,7]; G01N0033-68 [ICS,7]
AT 315228
                IPCI
                IPCR
                       G01N0033-53 [I,C*]; G01N0033-74 [I,C*]; G01N0033-53
                        [I,A]; G01N0033-74 [I,A]
```

ANSWER 23 OF 40 CAPLUS COPYRIGHT 2007 ACS on STN

```
ANSWER 2 OF 40 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
     1999:241472 BIOSIS
DN
     PREV199900241472
     An immunoluminometric assay for N-terminal pro-brain natriuretic
TI
     peptide: Development of a test for left ventricular dysfunction.
     Hughes, D.; Talwar, S.; Squire, I. B.; Davies, J. E.; Ng, L. L. [Reprint
AU
CS
     Department of Medicine and Therapeutics, Leicester Royal Infirmary,
     University of Leicester, Robert Kilpatrick Clinical Sciences Building,
     Leicester, LE2 7LX, UK
SO
     Clinical Science (London), (April, 1999) Vol. 96, No. 4, pp.
     373-380. print.
     CODEN: CSCIAE. ISSN: 0143-5221.
DT
     Article
     English
LA
ED
     Entered STN: 17 Jun 1999
     Last Updated on STN: 20 Aug 1999
AB
     Measurement of plasma levels of brain natriuretic peptide (
     BNP) has been used to assess left ventricular dysfunction and
     prognosis. Levels of the N-terminus of the precursor of BNP
     (NT-proBNP) have been reported to be elevated to a greater extent than
     BNP in left ventricular dysfunction. We have devised a
     non-radioactive sensitive and specific assay for NT-proBNP based on a
     competitive ligand binding principle. The chemiluminescent label
     4-(2-succinimidyl-oxycarbonylethyl)phenyl-10-methylacridinium
     9-carboxylate fluorosulphonate was used to label peptides representing
     domains in the middle and C-terminal sections of NT-proBNP. Assay of the
     C-terminal section of NT-proBNP (amino acids 65-76) in patients with
     proven left ventricular dysfunction (left ventricular wall motion index
     median 0.9 (range 0.3-1.4)) revealed elevated values (median 639 (386-911)
     fmol/ml) compared with normal controls (left ventricular wall motion index
     of 2 in all, NT-proBNP median 159 (120-245) fmol/ml, P < 0.001).
     Measurement of the middle section of NT-proBNP (amino acids 37-49) was not
     a discriminating test. It is thus possible to derivatize small peptides
     with a methyl acridinium label and preserve immunodetection with specific
     antibodies. Such methodology may allow non-radioactive
     immunoluminometric assays to be devised.
     Cardiovascular system - General and methods
     Biochemistry methods - General
                                      10050
     Endocrine - General
                           17002
     Biochemistry studies - General
                                      10060
     Major Concepts
IT
        Cardiovascular System (Transport and Circulation); Methods and
        Techniques
IT
     Diseases
        left ventricular dysfunction: heart disease
        Ventricular Dysfunction, Left (MeSH)
IT
     Chemicals & Biochemicals
         brain natriuretic peptide: plasma; N-terminal-pro-
       brain natriuretic peptide; 4-(2-succinimidyl-oxycarbonylethyl)
       phenyl-10-methylacridinium 9-carboxylate fluorosulfonate:
       chemiluminescent label
IT
    Methods & Equipment
        immunoluminometric assay: diagnostic method
RN
     114471-18-0 (brain natriuretic peptide)
     87198-89-8 (4-(2-succinimidyl-oxycarbonylethyl) phenyl-10-methylacridinium
     9-carboxylate fluorosulfonate)
     121128-24-3 (PRO-BRAIN NATRIURETIC PEPTIDE)
```

ANSWER 5 OF 55 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN

- AN 1997:510473 BIOSIS
- DN PREV199799809676
- TI A new, fast and reliable radioimmunoassay of brain natriuretic peptide in human plasma. Reference values in healthy subjects and in patients with different diseases.
- AU Jensen, K. T. [Reprint author]; Carstens, J.; Ivarsen, P.; Pedersen, E. B.
- CS Res. Lab. Nephrology and Hypertension, Aarhus Univ. Hosp., Aarhus Amtssyygehus, DK-8000 Aarhus C, Denmark
- SO Scandinavian Journal of Clinical and Laboratory Investigation, (1997) Vol. 57, No. 6, pp. 529-540.
  CODEN: SJCLAY. ISSN: 0036-5513.
- DT Article
- LA English
- ED Entered STN: 10 Dec 1997 Last Updated on STN: 10 Dec 1997

ANSWER 8 OF 55 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN

- AN 1996:191586 BIOSIS
- DN PREV199698747715
- TI Preparation of a monoclonal antibody against mouse brain natriuretic peptide (BNP) and tissue distribution of BNP in mice.
- AU Nakagawa, Masayo; Tanaka, Issei [Reprint author]; Suga, Shin-Ichi; Ogawa, Yoshihiro; Tamura, Naohisa; Goto, Masahisa; Sugawara, Akira; Yoshimasa, Takaaki; Itoh, Hiroshi; Mukoyama, Masashi; Nakao, Kazuwa
- CS Dep. Med. Clinical Sci., Kyoto Univ. Graduate Sch. Med., 54 Shogoin Kawahara-cho, Sakyo-ku, Kyoto 606, Japan
- SO Clinical and Experimental Pharmacology and Physiology, (1995) Vol. 22, No. SUPPL. 1, pp. S186-S187. ISSN: 0305-1870.
- DT Article
- LA English
- ED Entered STN: 2 May 1996 Last Updated on STN: 2 May 1996

ANSWER 16 OF 55 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN

- AN 1993:429763 BIOSIS
- DN PREV199396084388
- TI Ventricular expression of brain natriuretic peptide in hypertrophic cardiomyopathy.
- AU Hasegawa, Koji; Fujiwara, Hisayoshi [Reprint author]; Doyama, Kiyoshi; Miyamae, Masami; Fujiwara, Takako; Suga, Shinichi; Mukoyama, Masashi; Nakao, Kazuwa; Imura, Hiroo; Sasayama, Shigetake
- CS Third Div., Dep. Internal Med., Fac. Med., Kyoto Univ., 54 Kawara-cho Shogoin, Sakyo-ku, Kyoto 606, Japan
- SO Circulation, (1993) Vol. 88, No. 2, pp. 372-380. CODEN: CIRCAZ. ISSN: 0009-7322.
- DT Article
- LA English
- ED Entered STN: 22 Sep 1993 Last Updated on STN: 23 Sep 1993

ANSWER 5 OF 55 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN

- AN 1997:510473 BIOSIS
- DN PREV199799809676
- TI A new, fast and reliable radioimmunoassay of brain natriuretic peptide in human plasma. Reference values in healthy subjects and in patients with different diseases.
- AU Jensen, K. T. [Reprint author]; Carstens, J.; Ivarsen, P.; Pedersen, E. B.
- CS Res. Lab. Nephrology and Hypertension, Aarhus Univ. Hosp., Aarhus Amtssyygehus, DK-8000 Aarhus C, Denmark
- SO Scandinavian Journal of Clinical and Laboratory Investigation, (1997) Vol. 57, No. 6, pp. 529-540.
  CODEN: SJCLAY. ISSN: 0036-5513.
- DT Article
- LA English
- ED Entered STN: 10 Dec 1997
  Last Updated on STN: 10 Dec 1997

10/714,078 wpdated Lycook 1/12/04

## d his

(FILE 'HOME' ENTERED AT 14:35:33 ON 12 JAN 2007)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, JAPIO' ENTERED AT 14:35:48 ON 12 JAN 2007

	JAN 2007	
L1	11945	S BNP
L2	1787	S L1 AND HEMO?
L3	49	S L2 AND THROMBO?
L4	29	DUPLICATE REMOVE L3 (20 DUPLICATES REMOVED)
L5	1	S L4 AND ANTIBOD?
L6	28	S L4 NOT L5

ANSWER 12 OF 28 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN 1998:89478 BIOSIS ANDN PREV199800089478 Plasma brain natriuretic peptide levels increase in proportion to the TΙ extent to right ventricular dysfunction in pulmonary hypertension. Nagaya, Noritoshi; Nishikimi, Toshio [Reprint author]; Okano, Yoshiaki; AU Uematsu, Masaaki; Satoh, Toru; Kyotani, Shingo; Kuribayashi, Sachio; Hamada, Seiki; Kakishita, Mikio; Nakanishi, Norifumi; Takamiya, Makoto; Kunieda, Takeyoshi; Matsuo, Hisayuki; Kanagawa, Kenji Dep. Hypertension, Natl. Cardiovasc. Cent., 5-7-1 Fujishirodai, Suita, CS Osaka 565, Japan SO Journal of the American College of Cardiology, (Jan., 1998) Vol. 31, No. 1, pp. 202-208. print. CODEN: JACCDI. ISSN: 0735-1097. DT Article English LA ED Entered STN: 25 Feb 1998 Last Updated on STN: 25 Feb 1998 Objectives. This study sought to investigate the influence of right AB ventricular (RV) hemodynamic variables and function on the secretion of brain natriuretic peptide (BNP) in patients with isolated RV overload. Background. Plasma BNP is known to increase in proportion to the degree of left ventricular (LV) overload. However, whether BNP secretion is also regulated in the presence of RV overload remains unknown. Methods. Plasma BNP and atrial natriuretic peptide (ANP) levels in the pulmonary artery were measured in 44 patients with RV overload: 18 with RV volume overload (RVVO) due to atrial septal defect and 26 with RV pressure overload (RVPO) due to primary or thromboembolic pulmonary hypertension. Right heart catheterization was performed in all patients. RV and LV ejection fraction, myocardial mass and volume of the four chambers were determined by using electron beam computed tomography. Results. Although both plasma BNP and ANP levels were significantly elevated in patients with RV overload compared with values in control subjects, plasma BNP and the BNP/ANP ratio were significantly higher in patients with RVPO than with RVVO (BNP 294 +- 72 vs. 48 +- 14 pg/ml; BNP/ANP 1.6 +- 0.2 vs. 0.8 +- 0.2, both p < 0.05). Plasma BNP correlated positively with mean pulmonary artery pressure (r = 0.73), total pulmonary resistance (r = 0.79), mean right atrial pressure (r = 0.79), RV end-diastolic pressure (r = 0.76) and RV myocardial mass (r = 0.71); it correlated negatively with cardiac output (r = -0.33) and RV ejection fraction (r = -0.71). Plasma BNP significantly decreased from 315 +- 120 to 144 +- 54 pg/ml with long-term vasodilator therapy (total pulmonary resistance decreased from 23 +- 4 to 15 +- 3 Wood U). Conclusions. Plasma BNP increases in proportion to the extent of RV dysfunction in pulmonary hypertension. CC Cardiovascular system - General and methods 14501 Pathology - Therapy 12512 Endocrine - General 17002 Pharmacology - General 22002 IT Major Concepts Cardiovascular Medicine (Human Medicine, Medical Sciences) IT Diseases pulmonary hypertension: vascular disease Hypertension, Pulmonary (MeSH) IT Chemicals & Biochemicals atrial natriuretic peptide: secretion, plasma; brain natriuretic peptide: secretion, plasma; vasodilators IT Miscellaneous Descriptors cardiac output; mean right atrial pressure; pulmonary artery pressure; right ventricular ejection fraction; right ventricular end-diastolic pressure; right ventricular myocardial mass; right ventricular pressure overload; right ventricular volume overload; total pulmonary resistance

ANSWER 12 OF 28 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN AN 1998:89478 BIOSIS DN PREV199800089478 Plasma brain natriuretic peptide levels increase in proportion to the TI extent to right ventricular dysfunction in pulmonary hypertension. Nagaya, Noritoshi; Nishikimi, Toshio [Reprint author]; Okano, Yoshiaki; Uematsu, Masaaki; Satoh, Toru; Kyotani, Shingo; Kuribayashi, Sachio; Hamada, Seiki; Kakishita, Mikio; Nakanishi, Norifumi; Takamiya, Makoto; Kunieda, Takeyoshi; Matsuo, Hisayuki; Kanagawa, Kenji Dep. Hypertension, Natl. Cardiovasc. Cent., 5-7-1 Fujishirodai, Suita, CS Osaka 565, Japan Journal of the American College of Cardiology, (Jan., 1998) Vol. 31, No. SO 1, pp. 202-208. print. CODEN: JACCDI. ISSN: 0735-1097. DT Article LΑ English ED Entered STN: 25 Feb 1998 Last Updated on STN: 25 Feb 1998 AB Objectives. This study sought to investigate the influence of right ventricular (RV) hemodynamic variables and function on the secretion of brain natriuretic peptide (BNP) in patients with isolated RV overload. Background. Plasma BNP is known to increase in proportion to the degree of left ventricular (LV) overload. However, whether BNP secretion is also regulated in the presence of RV overload remains unknown. Methods. Plasma BNP and atrial natriuretic peptide (ANP) levels in the pulmonary artery were measured in 44 patients with RV overload: 18 with RV volume overload (RVVO) due to atrial septal defect and 26 with RV pressure overload (RVPO) due to primary or thromboembolic pulmonary hypertension. Right heart catheterization was performed in all patients. RV and LV ejection fraction, myocardial mass and volume of the four chambers were determined by using electron beam computed tomography. Results. Although both plasma BNP and ANP levels were significantly elevated in patients with RV overload compared with values in control subjects, plasma BNP and the BNP/ANP ratio were significantly higher in patients with RVPO than with RVVO (BNP 294 +- 72 vs. 48 +- 14 pq/ml; BNP/ANP 1.6 +- 0.2 vs. 0.8 +- 0.2, both p < 0.05). Plasma BNP correlated positively with mean pulmonary artery pressure (r = 0.73), total pulmonary resistance (r = 0.79), mean right atrial pressure (r = 0.79), RV end-diastolic pressure (r = 0.76) and RV myocardial mass (r = 0.71); it correlated negatively with cardiac output (r = -0.33) and RV ejection fraction (r = -0.71). Plasma BNP significantly decreased from 315 +- 120 to 144 +- 54 pg/ml with long-term vasodilator therapy (total pulmonary resistance decreased from 23 +- 4 to 15 +- 3 Wood U). Conclusions. Plasma BNP increases in proportion to the extent of RV dysfunction in pulmonary hypertension. Cardiovascular system - General and methods CC 14501 Pathology - Therapy 12512 Endocrine - General 17002 22002 Pharmacology - General IT Major Concepts Cardiovascular Medicine (Human Medicine, Medical Sciences) IT Diseases pulmonary hypertension: vascular disease Hypertension, Pulmonary (MeSH) IT Chemicals & Biochemicals atrial natriuretic peptide: secretion, plasma; brain natriuretic peptide: secretion, plasma; vasodilators IT Miscellaneous Descriptors cardiac output; mean right atrial pressure; pulmonary artery pressure; right ventricular ejection fraction; right ventricular end-diastolic pressure; right ventricular myocardial mass; right ventricular pressure overload; right ventricular volume overload; total pulmonary resistance ORGN Classifier

Hominidae 86215

Super Taxa

Primates; Mammalia; Vertebrata; Chordata; Animalia

Organism Name
human

Taxa Notes

Animals, Chordates, Humans, Mammals, Primates, Vertebrates

RN 85637-73-6 (atrial natriuretic peptide)

114471-18-0 (brain natriuretic peptide)

ORGN Classifier

Hominidae 86215

Super Taxa

Primates; Mammalia; Vertebrata; Chordata; Animalia

Organism Name
human

Taxa Notes
Animals, Chordates, Humans, Mammals, Primates, Vertebrates

RN 85637-73-6 (atrial natriuretic peptide)

114471-18-0 (brain natriuretic peptide)

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FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, JAPIO' ENTERED AT 14:35:48 ON 12 JAN 2007

	JAN 2007	
L1	11945	S BNP
L2	1787	S L1 AND HEMO?
L3	49	S L2 AND THROMBO?
L4	29	DUPLICATE REMOVE L3 (20 DUPLICATES REMOVED)
L5	1	S L4 AND ANTIBOD?
L6	28	S L4 NOT L5

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ANSWER 2 OF 40 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
AN
     1999:241472 BIOSIS
DN
     PREV199900241472
     An immunoluminometric assay for N-terminal pro-brain natriuretic
ΤI
     peptide: Development of a test for left ventricular dysfunction.
ΑU
     Hughes, D.; Talwar, S.; Squire, I. B.; Davies, J. E.; Ng, L. L. [Reprint
CS
     Department of Medicine and Therapeutics, Leicester Royal Infirmary,
     University of Leicester, Robert Kilpatrick Clinical Sciences Building,
     Leicester, LE2 7LX, UK
     Clinical Science (London), (April, 1999) Vol. 96, No. 4, pp.
SO
     373-380. print.
     CODEN: CSCIAE. ISSN: 0143-5221.
DT
     Article
LΑ
     English
ED
     Entered STN: 17 Jun 1999
     Last Updated on STN: 20 Aug 1999
     Measurement of plasma levels of brain natriuretic peptide (
AR
     BNP) has been used to assess left ventricular dysfunction and
     prognosis. Levels of the N-terminus of the precursor of BNP
     (NT-proBNP) have been reported to be elevated to a greater extent than
     BNP in left ventricular dysfunction. We have devised a
     non-radioactive sensitive and specific assay for NT-proBNP based on a
     competitive ligand binding principle. The chemiluminescent label
     4-(2-succinimidyl-oxycarbonylethyl)phenyl-10-methylacridinium
     9-carboxylate fluorosulphonate was used to label peptides representing
     domains in the middle and C-terminal sections of NT-proBNP. Assay of the
     C-terminal section of NT-proBNP (amino acids 65-76) in patients with
     proven left ventricular dysfunction (left ventricular wall motion index
     median 0.9 (range 0.3-1.4)) revealed elevated values (median 639 (386-911)
     fmol/ml) compared with normal controls (left ventricular wall motion index
     of 2 in all, NT-proBNP median 159 (120-245) fmol/ml, P < 0.001).
     Measurement of the middle section of NT-proBNP (amino acids 37-49) was not
     a discriminating test. It is thus possible to derivatize small peptides
     with a methyl acridinium label and preserve immunodetection with specific
     antibodies. Such methodology may allow non-radioactive
     immunoluminometric assays to be devised.
     Cardiovascular system - General and methods
     Biochemistry methods - General
                                      10050
     Endocrine - General
                           17002
     Biochemistry studies - General
                                      10060
IT
     Major Concepts
        Cardiovascular System (Transport and Circulation); Methods and
        Techniques
IT
     Diseases
        left ventricular dysfunction: heart disease
        Ventricular Dysfunction, Left (MeSH)
IT
     Chemicals & Biochemicals
          brain natriuretic peptide: plasma; N-terminal-pro-
        brain natriuretic peptide; 4-(2-succinimidyl-oxycarbonylethyl)
        phenyl-10-methylacridinium 9-carboxylate fluorosulfonate:
        chemiluminescent label
IT
     Methods & Equipment
        immunoluminometric assay: diagnostic method
RN
     114471-18-0 (brain natriuretic peptide)
     87198-89-8 (4-(2-succinimidyl-oxycarbonylethyl) phenyl-10-methylacridinium
```

9-carboxylate fluorosulfonate)

121128-24-3 (PRO-BRAIN NATRIURETIC PEPTIDE)

```
ANSWER 30 OF 40 EMBASE COPYRIGHT (c) 2007 Elsevier B.V. All rights
     reserved on STN
AN
     1998012811 EMBASE
     Biochemical detection of left-ventricular systolic dysfunction.
TТ
     McDonagh T.A.; Robb S.D.; Murdoch D.R.; Morton J.J.; Ford I.; Morrison
ΑU
     C.E.; Tunstall-Pedoe H.; McMurray J.J.V.; Dargie H.J.
CS
     Dr. T.A. McDonagh, Cardiology Department, Western Infirmary, Glasgow G11
     6NT, United Kingdom
SO
     Lancet, (3 Jan 1998) Vol. 351, No. 9095, pp. 9-13. .
     Refs: 27
     ISSN: 0140-6736 CODEN: LANCAO
CY
     United Kingdom
DT
     Journal; Article
FS
             Internal Medicine
     006
             Cardiovascular Diseases and Cardiovascular Surgery
     018
     029
             Clinical Biochemistry
     036
             Health Policy, Economics and Management
LΑ
     English
SL
     English
     Entered STN: 22 Jan 1998
ED
     Last Updated on STN: 22 Jan 1998
AΒ
     Background: In previous studies on the use of natriuretic peptides to
     detect left-ventricular systolic dysfunction, a higher rate of cardiac
     disorders in the control groups than in the study groups could have led to
           We investigated the effectiveness of plasma N-terminal atrial
     natriuretic peptide (NT-ANP) and brain natriuretic peptide (
     BNP) concentrations to show left-ventricular systolic dysfunction
     in a random sample of the general population. Methods: We randomly
     selected 2000 participants aged 25-74 years from family physicians' lists
     in Glasgow, UK. We sent all participants questionnaires. 1653 respondents
     underwent echocardiography and electrocardiography. We took a
     left-ventricular ejection fraction of 30% or less to show left-ventricular
     systolic dysfunction. NT-ANP and BNP were measured in plasma by
     RIAs. Findings: 1252 participants had analysable electrocardiograms and
     echocardiograms, completed questionnaires, and available blood samples.
    Median concentrations of NT-ANP and BNP were significantly
    higher in participants with left-ventricular systolic dysfunction (2.8
    ng/mL [IQR 1.8-4.6] and 24.0 pg/mL [18.0-33.0]) than in those without (1.3
    ng/mL [0.9-1.8] and 7.7 pg/mL [3.4-13.0]; each p < 0.001). Among
    participants with left-ventricular systolic dysfunction, both symptomatic
    and asymptomatic subgroups had raised NT-ANP and BNP
     concentrations. A BNP concentration of 17.9 pg/mL or more gave
    a sensitivity of 77% and specificity of 87% in all participants, and 92%
    and 72% in participants aged 55 years or older. Interpretation:
    Measurement of BNP could be a cost-effective method of screening
     for left-ventricular systolic dysfunction in the general population,
    especially if its use were targeted to individuals at high risk.
    Medical Descriptors:
     *heart left ventricle failure: DI, diagnosis
    biochemistry
    hormone blood level
    united kingdom
    echocardiography
    electrocardiography
    heart ejection fraction
    radioimmunoassay
    questionnaire
    cost effectiveness analysis
    high risk population
    human
    male
    female
    major clinical study
    controlled study
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ANSWER 3 OF 5 EMBASE COPYRIGHT (c) 2007 Elsevier B.V. All rights
     reserved on STN
ΑN
     81094356 EMBASE
DN
     1981094356
     Computerized axial tomography in the detection of brain damage. 2.
ΤI
     Epilepsy, migraine, and general medical disorders.
     Cala L.A.; Mastaglia F.L.
AU
     Dept. Diagn. Radiol., Sir Charles Gairdner Hosp., Queen Elizabeth Med.
CS
     Cent., Perth, Australia
     Medical Journal of Australia, (1980) Vol. 2, No. 11, pp. 616-620. .
so
     CODEN: MJAUAJ
CY
     Australia
DT
     Journal
FS
     014
             Radiology
     008
             Neurology and Neurosurgery
     027
             Biophysics, Bioengineering and Medical Instrumentation
     050
             Epilepsy
LA
     English
     Entered STN: 9 Dec 1991
     Last Updated on STN: 9 Dec 1991
     The cranial computerized axial tomography (CAT) findings in groups of
AB
     patients with epilepsy, migraine, hypertension, and other general medical
     disorders have been reviewed to assess the frequency and
     patterns of focal and diffuse brain damage. In addition to demonstrating
     focal lesions in a proportion of patients with seizures and in patients
     presenting with a stroke, the CAT scan
     showed a premature degree of cerebral atrophy in an appreciable proportion
     of patients with long-standing epilepsy, hypertension and diabetes, and in
     some patients with migraine, valvular and ischaemic heart disease, chronic
     obstructive airways disease, and chronic renal failure. The value of CAT
     as a means of screening for brain damage in groups of individuals at risk
     is discussed.
     Medical Descriptors:
     *brain atrophy
     *computer assisted tomography
     *epilepsy
     *hydrocephalus
     *hypertension
     *migraine
     central nervous system
     diagnosis
     major clinical study
     computer analysis
     cardiovascular system
```